

WHAT IS CLAIMED IS:

1. A method of creating a chemical compound library comprising:  
selecting compounds having a molecular weight of no greater  
5 than about 350 grams/mole; and  
selecting compounds having a solubility in deuterated water of at  
least about 1 mM at room temperature.
2. The method of claim 1 wherein a majority of the compounds in the  
10 chemical compound library have a molecular weight of no greater than  
about 350 grams/mole and a solubility in deuterated water of at least  
about 1 mM at room temperature.
3. The method of claim 2 wherein all of the compounds in the chemical  
15 compound library have a molecular weight of no greater than about 350  
grams/mole and a solubility in deuterated water of at least about 1 mM at  
room temperature.
4. The method of claim 1 wherein the compounds selected have a molecular  
20 weight of no greater than about 325 grams/mole.
5. The method of claim 4 wherein the compounds selected have a molecular  
weight of less than about 325 grams/mole.
- 25 6. A chemical compound library comprising compounds having a molecular  
weight of no greater than about 350 grams/mole and a solubility in  
deuterated water of at least about 1 mM at room temperature.
7. The library of claim 6 wherein a majority of the compounds have a  
30 molecular weight of no greater than about 350 grams/mole and a  
solubility in deuterated water of at least about 1 mM at room temperature.
8. The library of claim 7 wherein all of the compounds have a molecular

weight of no greater than about 350 grams/mole and a solubility in deuterated water of at least about 1 mM at room temperature.

9. The library of claim 6 wherein the compounds have a molecular weight  
5 of no greater than about 325 grams/mole.
10. The library of claim 9 wherein the compounds have a molecular weight  
of less than about 325 grams/mole.
- 10 11. A method of identifying a lead chemical template, the method  
comprising:  
selecting compounds having a molecular weight of no greater  
than about 350 grams/mole and a solubility in deuterated water of at least  
about 1 mM at room temperature to create a chemical compound library;  
15 identifying at least one compound from the library that functions  
as a ligand to a target molecule having a dissociation constant of at least  
about 100  $\mu$ M; and  
using the ligand to identify a lead chemical template.
- 20 12. The method of claim 11 wherein a majority of the compounds in the  
chemical compound library have a molecular weight of no greater than  
about 350 grams/mole and a solubility in deuterated water of at least  
about 1 mM at room temperature.
- 25 13. The method of claim 12 wherein all of the compounds in the chemical  
compound library have a molecular weight of no greater than about 350  
grams/mole and a solubility in deuterated water of at least about 1 mM at  
room temperature.
- 30 14. The method of claim 11 wherein the compounds selected for the library  
have a molecular weight of no greater than about 325 grams/mole.
15. The method of claim 14 wherein the compounds selected for the library

have a molecular weight of less than about 325 grams/mole.

16. The method of claim 11 wherein the dissociation constant of a lead chemical template to the target molecule is at least about 1  $\mu\text{M}$ .

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17. The method of claim 11 wherein the target molecule is a protein.

18. A method of identifying a compound that binds to a target molecule, the method comprising:

10 providing a plurality of mixtures of test compounds, each mixture being in a sample reservoir;

introducing a target molecule into each of the sample reservoirs to provide a plurality of test samples;

15 providing a nuclear magnetic resonance spectrometer equipped with a flow-injection probe;

transferring each test sample from the sample reservoir into the flow-injection probe;

collecting a relaxation-edited nuclear magnetic resonance spectrum on each sample in each reservoir; and

20 comparing the spectra of each sample to the spectra taken under the same conditions in the absence of the target molecule to identify compounds that bind to the target molecule;

wherein the concentration of target molecule and each compound in each sample is no greater than about 100  $\mu\text{M}$ .

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19. The method of claim 18 wherein each mixture is in a sample reservoir of a multiwell sample holder.

20. The method of claim 19 wherein the multiwell sample holder is a 96-well  
30 microtiter plate.

21. The method of claim 18 wherein each test compound has a solubility in deuterated water of at least about 1 mM at room temperature.
22. The method of claim 18 wherein each test compound has a molecular weight of no greater than about 350 grams/mole.
23. The method of claim 18 wherein collecting a relaxation-edited nuclear magnetic resonance spectrum comprises collecting a 1D relaxation-edited nuclear magnetic resonance spectrum.
24. The method of claim 23 wherein collecting a 1D relaxation-edited nuclear magnetic resonance spectrum comprises collecting a 1D relaxation-edited  $^1\text{H}$  nuclear magnetic resonance spectrum.
25. The method of claim 18 wherein the mixture of compounds comprises at least about 3 compounds, each having at least one distinguishable resonance in a 1D NMR spectrum of the mixture.
26. The method of claim 25 wherein the mixture of compounds comprises at least about 6 compounds.
27. The method of claim 25 wherein the ratio of target molecule to each test compound in each sample reservoir is about 1:1.
28. The method of claim 18 wherein the concentration of target molecule and each compound in each sample is no greater than about 50  $\mu\text{M}$ .
29. The method of claim 18 wherein the dissociation constant of a compound that binds to the target molecule is at least about 100  $\mu\text{M}$ .
30. The method of claim 18 wherein the target molecule is a protein.
31. A method of identifying a compound that binds to a target molecule, the

method comprising:

providing a plurality of mixtures of test compounds, each mixture being in a sample reservoir;

5 introducing a target molecule into each of the sample reservoirs to provide a plurality of test samples;

providing a nuclear magnetic resonance spectrometer equipped with a flow-injection probe;

10 transferring each test sample from the sample reservoir into the flow-injection probe;

collecting a WaterLOGSY nuclear magnetic resonance spectrum on each sample in each reservoir; and

15 analyzing the spectra of each sample to distinguish binding compounds from nonbinding compounds by virtue of the opposite sign of their water-ligand NOEs.

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32. The method of claim 31 wherein the concentration of target molecule is no greater than about 10  $\mu\text{M}$ .

20 33. The method of claim 32 wherein the concentration of target molecule is no greater than about 1  $\mu\text{M}$ .

34. The method of claim 31 wherein the concentration of each compound in each sample is no greater than about 100  $\mu\text{M}$ .

25 35. The method of claim 31 wherein each test compound has a solubility in deuterated water of at least about 1 mM at room temperature.

36. The method of claim 31 wherein each mixture is in a sample reservoir of a multiwell sample holder.

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37. The method of claim 36 wherein the multiwell sample holder is a 96-well microtiter plate.

38. The method of claim 31 wherein each test compound has a molecular weight of no greater than about 350 grams/mole.
- 5 39. The method of claim 38 wherein each test compound has a molecular weight of no greater than about 325 grams/mole.
40. The method of claim 31 wherein collecting a WaterLOGSY nuclear magnetic resonance spectrum comprises collecting a 1D WaterLOGSY  
10 nuclear magnetic resonance spectrum.
41. The method of claim 31 wherein the mixture of compounds comprises at least about 3 compounds, each having at least one distinguishable resonance in a 1D NMR spectrum of the mixture.  
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42. The method of claim 41 wherein the mixture of compounds comprises at least about 6 compounds.
43. The method of claim 31 wherein the ratio of target molecule to each test  
20 compound in each sample reservoir is about 100:1 to about 10:1.
44. The method of claim 31 wherein the dissociation constant of a compound that binds to the target molecule is at least about 100  $\mu$ M.
- 25 45. The method of claim 31 wherein the target molecule is a protein.
46. A method of identifying a protein function, the method comprising:  
providing a plurality of mixtures of test compounds consisting of  
known inhibitors, cofactors, and substrates of known proteins, each  
30 mixture being in a sample reservoir and containing a plurality of test  
compounds;  
introducing a target molecule into each of the sample reservoirs to  
provide a plurality of test samples;

providing a nuclear magnetic resonance spectrometer equipped  
with a flow-injection probe;

transferring each test sample from the sample reservoir into the  
flow-injection probe;

5           collecting a WaterLOGSY nuclear magnetic resonance spectrum  
on each sample in each reservoir;

          comparing the spectra of each sample to the spectra taken under  
the same conditions in the absence of the target molecule to identify  
compounds that bind to the target molecule, wherein the concentration of  
10           target molecule and each compound in each sample is no greater than  
about 5  $\mu$ M and 125  $\mu$ M, respectively; and

          determining a function of the target molecule based upon the test  
compounds that bind to the target molecule.

PATENT APPLICATION